

1/7

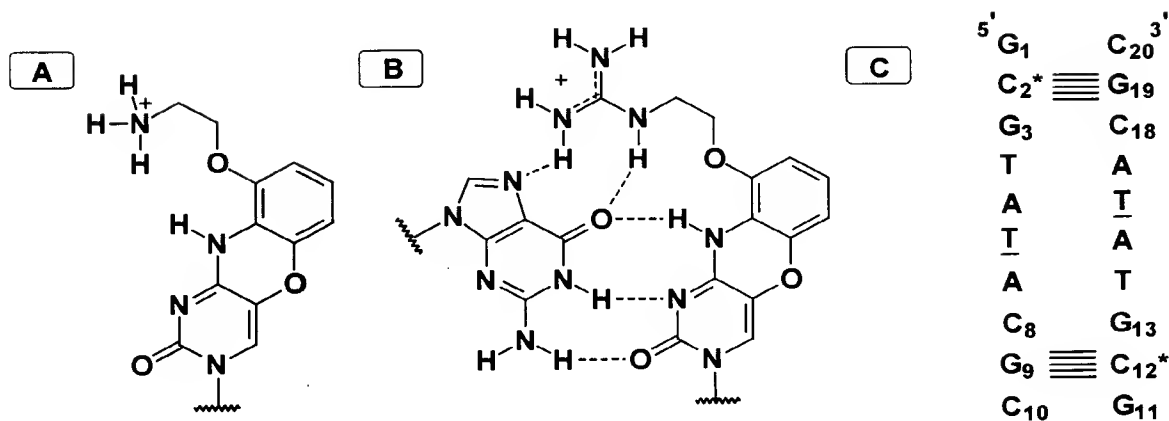


Figure 1. Structure of the tricyclic cytosine analog G-clamp⁵ (A), of its extended analog guanyl G-clamp hybridized to complementary guanosine (B), and of the palindromic decamer duplex crystallized for this study (C). The five hydrogen bonds formed between C* and G are indicated by horizontal lines (C* = guanyl G-clamp, T = 2'-O-MOE-T).

2/7

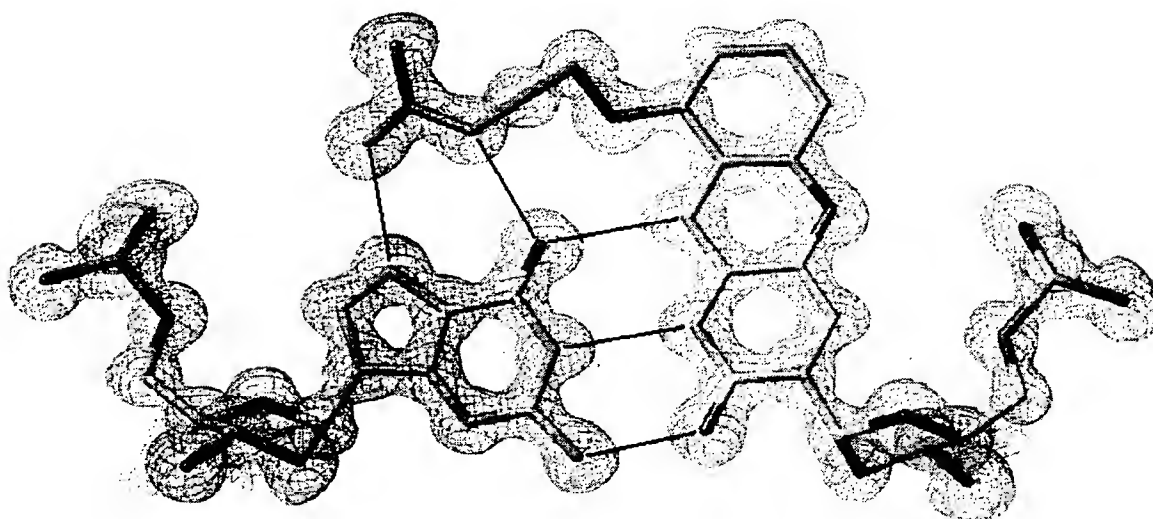


Figure 2. Fourier ($2F_o - F_o$) sum electron density map (contoured at 1.25σ) around C12* and G9 confirming formation of five hydrogen bonds (indicated by thin solid lines with distances shown in Å).

3/7

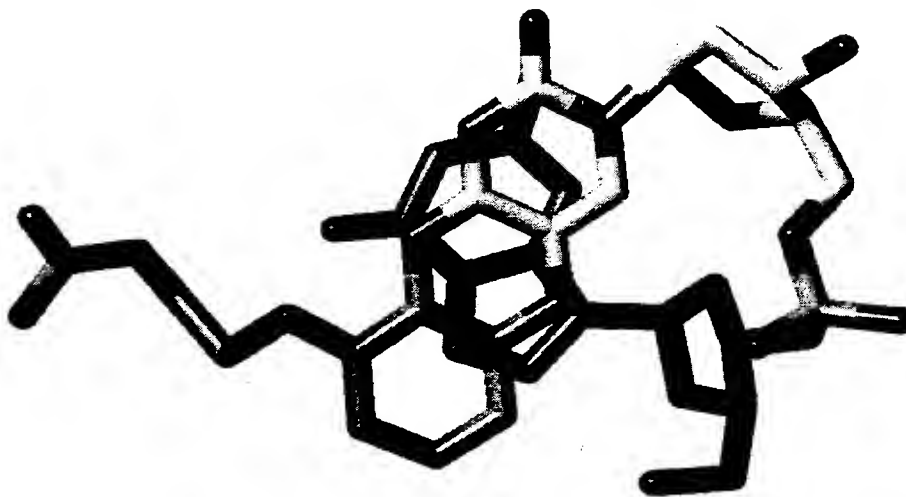


Figure 3. Stacking between G1 and C2*, viewed approximately along the vertical to the phenoxazine rings. Carbon atoms of G1 are shown in magenta, carbon atoms of the cytosine core of C2* are shown in yellow and the remainder of the carbons are in green.

4/7

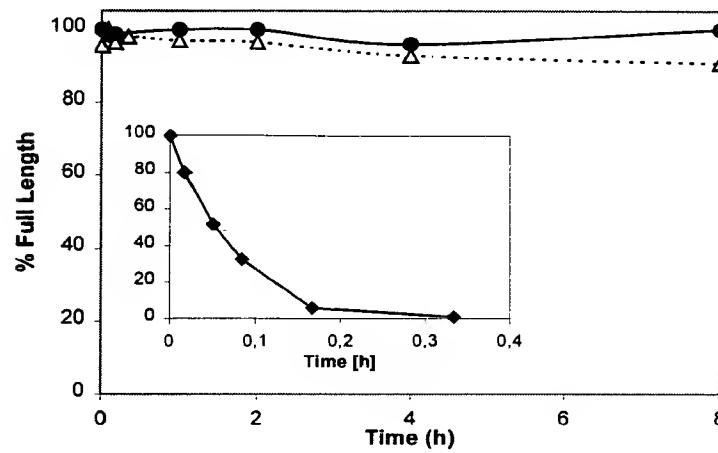


Figure 4

Figure 4. Degradation of ONs 157 (open triangles) and 158 (closed circles) as a function of incubation time and compared to an unmodified control ON 159 (closed diamonds, insert) determined by CGE analysis.

5/7

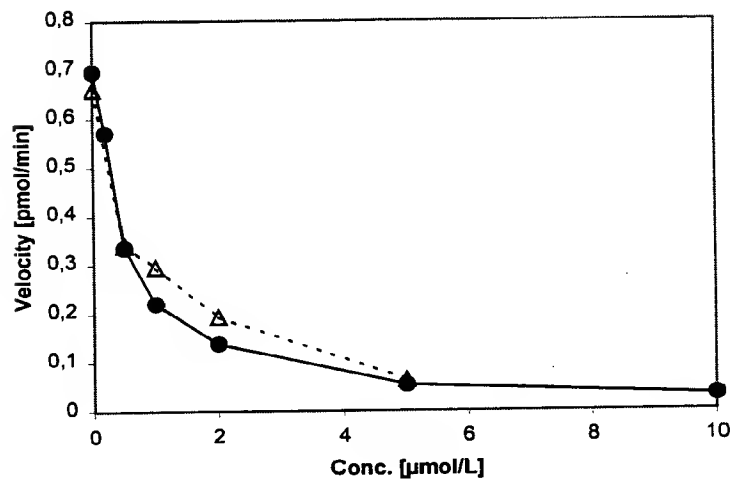


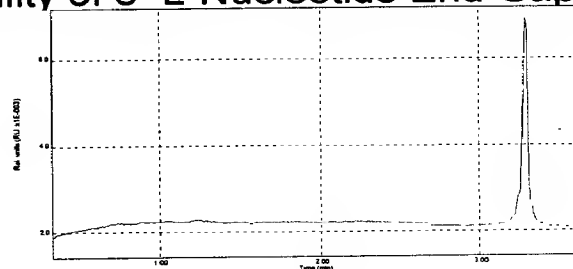
Figure 5.

Figure 5. Velocity of the enzymatic reaction: hydrolysis of ON 159 with BIPD as a function of the concentration of co-incubated ON 157 (open triangles) and ON 158 (closed circles).

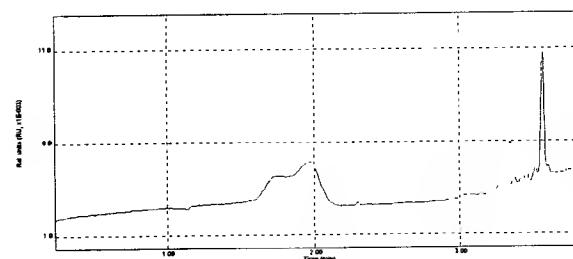
6/7

Stability of 3'-L-Nucleotide End-Capped

BalbC
mice
25 mg/kg
dose
i.v.



Starting Material



Liver @ 1hr

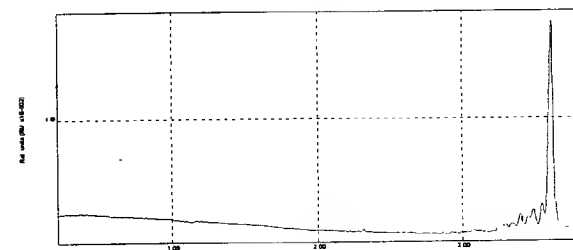
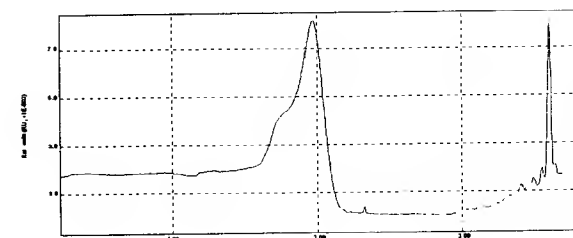
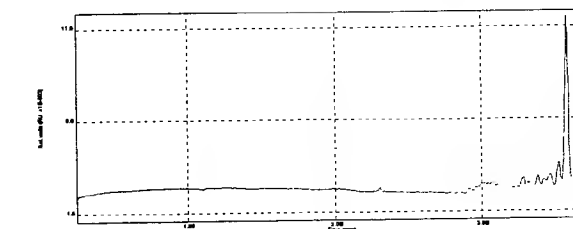
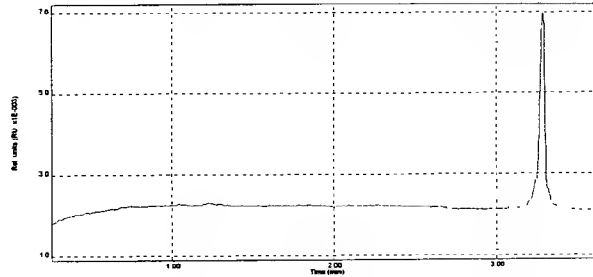
Kidney @ 1
hrSpleen @ 1
hrLung @
1 hr

Figure 6

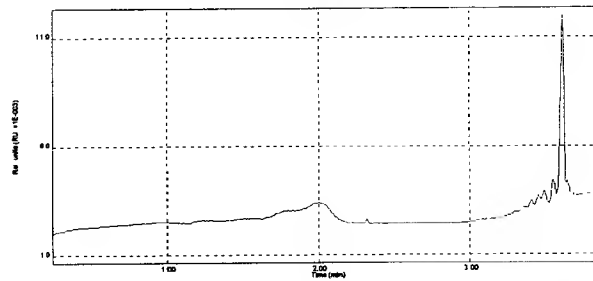
7/7

Stability of 3'-L-Nucleotide End-Capped

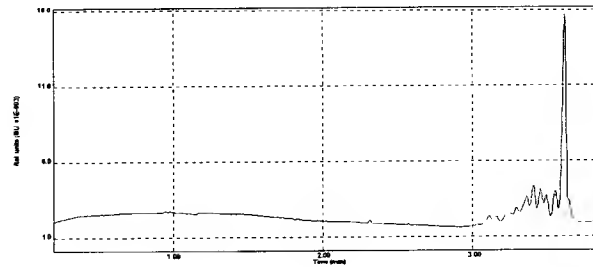
Starting Material



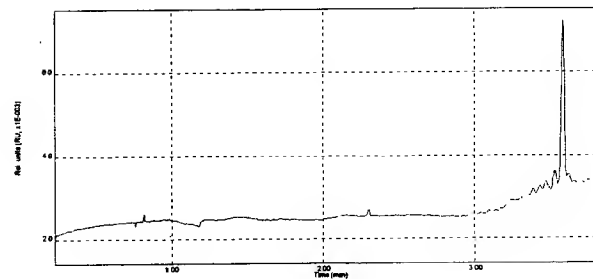
Liver @ 24 hr



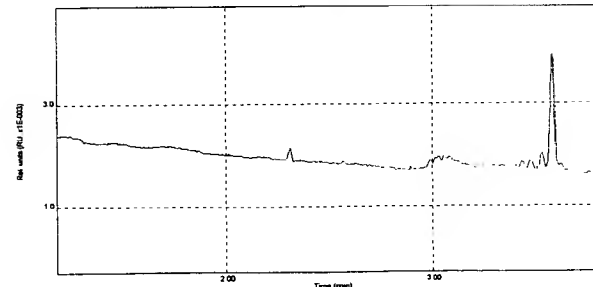
Kidney @ 24 hr



Spleen @ 24 hr



Lung @ 24 hr



BalbC
 mice
 25 mg/kg
 dose
 i.v.

Figure 7